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OIPE

RAW SEQUENCE LISTING DATE: 07/19/2001 PATENT APPLICATION: US/09/788,188 TIME: 12:09:47

Input Set : A:\41673204.app

Output Set: N:\CRF3\07192001\I788188.raw

```
3 <110> APPLICANT: TUSZYNSKI, MARK
       BLESCH, ARMIN
 6 <120> TITLE OF INVENTION: MUTANT PRO-NEUROTROPHIN WITH IMPROVED ACTIVITY
 8 <130> FILE REFERENCE: 041673/2045
10 <140> CURRENT APPLICATION NUMBER: 09/788,188
11 <141> CURRENT FILING DATE: 2001-02-16
13 <160> NUMBER OF SEQ ID NOS: 16
                                                        ENTERED
15 <170> SOFTWARE: PatentIn Ver. 2.1
17 <210> SEQ ID NO: 1
18 <211> LENGTH: 241
19 <212> TYPE: PRT
20 <213> ORGANISM: Homo sapiens
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                                        10
26 Gln Ala Glu Pro His Ser Glu Ser Asn Val Pro Ala Gly His Thr Ile
                                    25
29 Pro Gln Val His Trp Thr Lys Leu Gln His Ser Leu Asp Thr Ala Leu
            35
32 Arg Arg Ala Arg Ser Ala Pro Ala Ala Ala Ile Ala Ala Arg Val Ala
                            55
35 Gly Gln Thr Arg Asn Ile Thr Val Asp Pro Arg Leu Phe Lys Lys Arg
                        70
                                            75
38 Arg Leu Arg Ser Pro Arg Val Leu Phe Ser Thr Gln Pro Pro Arg Glu
                    85
                                        90
41 Ala Ala Asp Thr Gln Asp Leu Asp Phe Glu Val Gly Gly Ala Ala Pro
               100
                                   105
44 Phe Asn Arg Thr His Arg Ser Lys Arg Ser Ser Ser His Pro Ile Phe
          115
                               120
                                                   125
47 His Arg Gly Glu Phe Ser Val Cys Asp Ser Val Ser Val Trp Val Gly
                           135
50 Asp Lys Thr Thr Ala Thr Asp Ile Lys Gly Lys Glu Val Met Val Leu
                       150
53 Gly Glu Val Asn Ile Asn Asn Ser Val Phe Lys Gln Tyr Phe Phe Glu
                                       170
56 Thr Lys Cys Arg Asp Pro Asn Pro Val Asp Ser Gly Cys Arg Gly Ile
               180
                                   185
59 Asp Ser Lys His Trp Asn Ser Tyr Cys Thr Thr Thr His Thr Phe Val
                               200
62 Lys Ala Leu Thr Met Asp Gly Lys Gln Ala Ala Trp Arg Phe Ile Arg
                           215
                                               220
65 Ile Asp Thr Ala Cys Val Cys Val Leu Ser Arg Lys Ala Val Arg Arg
66 225
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73 <211> LENGTH: 241
74 <212> TYPE: PRT
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Input Set : A:\41673204.app

Output Set: N:\CRF3\07192001\1788188.raw

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77 <220> FEATURE:
78 <223> OTHER INFORMATION: Description of Artificial Sequence: Mutant NGF
         pro-neurotrophin
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88 Pro Gln Val His Trp Thr Lys Leu Gln His Ser Leu Asp Thr Ala Leu
                                40
91 Arg Arg Ala Arg Ser Ala Pro Ala Ala Ala Ile Ala Ala Arg Val Ala
                            55
94 Gly Gln Thr Arg Asn Ile Thr Val Asp Pro Arg Leu Phe Lys Lys Arg
                        70
97 Arg Leu Arg Ser Pro Arg Val Leu Phe Ser Thr Gln Pro Pro Arg Glu
                    85
                                         90
100 Ala Ala Asp Thr Gln Asp Leu Asp Phe Glu Val Gly Gly Ala Ala Pro
                                    105
103 Phe Ser Arg Thr His Arg Ser Lys Arg Ser Ser Ser His Pro Ile Phe
            115
                                 120
106 His Arg Gly Glu Phe Ser Val Cys Asp Ser Val Ser Val Trp Val Gly
                            135
                                                 140
109 Asp Lys Thr Thr Ala Thr Asp Ile Lys Gly Lys Glu Val Met Val Leu
                        150
                                             155
112 Gly Glu Val Asn Ile Asn Asn Ser Val Phe Lys Gln Tyr Phe Phe Glu
                                        170
                    165
115 Thr Lys Cys Arg Asp Pro Asn Pro Val Asp Ser Gly Cys Arg Gly Ile
                180
                                    185
118 Asp Ser Lys His Trp Asn Ser Tyr Cys Thr Thr Thr His Thr Phe Val
           195
                                200
121 Lys Ala Leu Thr Met Asp Gly Lys Gln Ala Ala Trp Arg Phe Ile Arg
                            215
124 Ile Asp Thr Ala Cys Val Cys Val Leu Ser Arg Lys Ala Val Arg Arg
125 225
                        230
                                             235
127 Ala
131 <210> SEQ ID NO: 3
132 <211> LENGTH: 247
133 <212> TYPE: PRT
134 <213> ORGANISM: Homo sapiens
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140 Lys Ala Ala Pro Met Lys Glu Ala Asn Ile Arq Gly Gln Gly Leu
                                     25
143 Ala Tyr Pro Gly Val Arg Thr His Gly Thr Leu Glu Ser Val Asn Gly
146 Pro Lys Ala Gly Ser Arg Gly Leu Thr Ser Leu Ala Asp Thr Phe Glu
147
                             55
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RAW SEQUENCE LISTING DATE: 07/19/2001
PATENT APPLICATION: US/09/788,188 TIME: 12:09:47

Input Set : A:\41673204.app

Output Set: N:\CRF3\07192001\1788188.raw

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· 149 His Val Ile Glu Glu Leu Leu Asp Glu Asp Gln Lys Val Arg Pro Asn
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                                              75
 152 Glu Glu Asn Asn Lys Asp Ala Asp Leu Tyr Thr Ser Arg Val Met Leu
                      85
                                          90
 155 Ser Ser Gln Val Pro Leu Glu Pro Pro Leu Leu Phe Leu Leu Glu Glu
                                     105
 158 Tyr Lys Asn Tyr Leu Asp Ala Ala Asn Met Ser Met Arg Val Arg Arg
                                  120
 161 His Ser Asp Pro Ala Arg Arg Gly Glu Leu Ser Val Cys Asp Ser Ile
                             135
 164 Ser Glu Trp Val Thr Ala Ala Asp Lys Lys Thr Ala Val Asp Met Ser
                         150
                                             155
 167 Gly Gly Thr Val Thr Val Leu Glu Lys Val Pro Val Ser Lys Gly Gln
                     165
                                         170
 170 Leu Lys Gln Tyr Phe Tyr Glu Thr Lys Cys Asn Pro Met Gly Tyr Thr
                                     185
                 180
 173 Lys Glu Gly Cys Arg Gly Ile Asp Lys Arg His Trp Asn Ser Gln Cys
                                 200 -
 176 Arg Thr Thr Gln Ser Tyr Val Arg Ala Leu Thr Met Asp Ser Lys Lys
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                             215
 179 Arg Ile Gly Trp Arg Phe Ile Arg Ile Asp Thr Ser Cys Val Cys Thr
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 182 Leu Thr Ile Lys Arg Gly Arg
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 186 <210> SEO ID NO: 4
 187 <211> LENGTH: 247
 188 <212> TYPE: PRT
 189 <213> ORGANISM: Artificial Sequence
 191 <220> FEATURE:
192 <223> OTHER INFORMATION: Description of Artificial Sequence: Mutant BDNF
        pro-neurotrophin
195 <400> SEQUENCE: 4
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197 1
199 Lys Ala Ala Pro Met Lys Glu Ala Asn Ile Arg Gly Gln Gly Gly Leu
                  20
                                      25
202 Ala Tyr Pro Gly Val Arg Thr His Gly Thr Leu Glu Ser Val Asn Gly
205 Pro Lys Ala Gly Ser Arg Gly Leu Thr Ser Leu Ala Asp Thr Phe Glu
                              55
208 His Val Ile Glu Glu Leu Leu Asp Glu Asp Gln Lys Val Arg Pro Asn
209 65
                          70.
211 Glu Glu Asn Asn Lys Asp Ala Asp Leu Tyr Thr Ser Arg Val Met Leu
                     85
                                          90
214 Ser Ser Gln Val Pro Leu Glu Pro Pro Leu Leu Phe Leu Leu Glu Glu
                                     105
217 Tyr Lys Asn Tyr Leu Asp Ala Ala Ser Met Ser Met Arg Val Arg Arg
                                 120
220 His Ser Asp Pro Ala Arg Arg Gly Glu Leu Ser Val Cys Asp Ser Ile
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Input Set : A:\41673204.app

Output Set: N:\CRF3\07192001\1788188.raw

221		130					135					140				
	Ser	Glu	Trp	Val	Thr	Ala		asp	Lvs	Lvs	Thr		Val	Asp	Met	Ser
	145		-			150		•	•	-	155			-		160
226	Gly	Gly	Thr	Val	Thr	Val	Leu	Glu	Lys	Val	Pro	Val	Ser	Lys	Gly	Gln
227	_	-			165				-	170				-	175	
	Leu	Lys	Gln	Tvr	Phe	Tvr	Glu	Thr	Lvs		Asn	Pro	Met	Glv	Tvr	Thr
230		- 1		180		-1-			185	-1-				190	-1-	
	Lvs	Glu	Glv		Arg	Glv	Ile	Asp		Ara	His	Trp	Asn		Gln	Cvs
233	2		195	-1-	5	1		200	-1-	,			205			-1-
	Ara	Thr		Gln	Ser	Tvr	Val		Αla	Len	Thr	Met		Ser	Lvs	Lvs
236	5	210		<b>0</b>		-1-	215	9		200		220		001	210	272
	Arσ		Glv	Trn	Arg	Phe		Arσ	Tle	Asn	Thr		Cvs	Val	Cvs	Thr
	225		011		9	230		5			235	001	0,0	, 41	0,0	240
		Thr	Tle	Lvc	Arg		Δrα				233					240
242	пса	1 111	110	цуз	245	GLY	nig									
	<210	)> </td <td>EQ II</td> <td>NO.</td> <td></td>	EQ II	NO.												
			ENGTE													
			PE:		<i>,</i>											
					Homo		ai an	,								
			COME			Sal	oren:	•								
			-			Фиг	17 a 1	т1.	Dho	LOU	7 l a	Фттх	T 011	λωσ	C1 77	т10
	мес 1	ser	TTE	пеп	Phe 5	тут	vaı	тте	Pile		Ата	тут	ьец	Arg	15	TTE
		C1	) an	7 ~ ~	_	7	<i>a</i> 1 =	7 mm	C = ==	10	Dago	a1	7	0		3
255	GIII	СТА	ASII		Met	Asp				ьeu	Pro	GIU	ASP		Leu	ASII
	Com	Т о	т1.	20	T	r				3	<b>-</b> 1-	T	T	30	T	T
258	ser	ьеи	35	тте	Lys	ьеu	TTE		Ата	ASP	тте	ьeu	_	ASII	гуѕ	Leu
	Com	T		14-±	17- 1	3	T7 - 1	40	a1	3	m	<b>a</b> 1	45	m l	T	D
	ser		ĠŢIJ	мес	Val	ASP		ьуѕ	GIU	ASII	TAL		ser	THE	Leu	Pro
261	T	50	a1	7 1 n	D	3	55	D	a1	3	<b>a</b> 1	60	D		<b>*</b>	<b>Q</b>
		Ald	GIU	Ата	Pro		GIU	Pro	GIU	Arg	_	GIY	Pro	Ата	гàг	
264	65	Dha	<b>a</b> 1	D	17- 1	70		V-4	3	m \	75	T	T		a 1	80
	Ата	Pne	GIII	PLO	Val	тте	Ата	мет	Asp		GIU	Leu	Leu	Arg		GIn
267	7	7	m	<b>3</b>	85	D	3	77- 7	T	90			<b>a</b>	m1	95	<b>-</b>
	Arg	Arg	туг		Ser	Pro	Arg	vaı		Leu	ser	Asp	ser		Pro	Leu
270	<b>a</b> 1	D	D	100	T	·	T	<b></b>	105	•	<b></b>	** - 1	<b>a</b> 1	110	_	**. 3
272	GIU			PLO	Leu	Tyr	ьeu		GIU	ASP	Tyr	vaı	_	ser	Pro	vaı
	17n 1		115	7	m la	<b></b>	3	120	T	3	ш		125	TT 2 =	T	<b>a</b>
	vaı		ASn	Arg	Thr	ser		Arg	ьуs	Arg	Tyr		GIU	HIS	ьуs	ser
276	TT 4	130	<b>a</b> 1	<b>a</b> 1	Ф	<b>a</b>	135				<b>a</b> 1	140	_	_	1	
			GTA	GIU	Tyr	ser	vaı	cys					Leu	Trp		
	145		_								155			_,		160
	Asp	гÀг	ser	ser	Ala	тте	Asp	тте	Arg		HIS	GIn	Val	Thr		Leu
282	<b>a</b> 1	<b>a</b> 3	-1.	_	165	~ 1	_	_	_	170	_		_	_,	175	
	GIA	GIU	тте		Thr	GIŸ	Asn	Ser		Val	Lys	GIn	Tyr		Tyr	GLu
285	1.	_	_	180	~ 3		_	_	185	_	_		_	190		
	Tnr	Arg		гàг	Glu	Ala	Arg		٧al	rys	Asn	СТĀ		Arg	GTA	те
288	_	_	195	•	_	_	_	200	_	_			205		_	
	Asp		Lys	His	Trp	Asn		GIn	Cys	ГЛS	Thr		Gln	Thr	$\mathtt{Tyr}$	Val
291	_	210	_		_		215	_	_			220		_	_	
293	Arg	Ala	Leu	Thr	Ser	Glu	Asn	Asn	Lys	Leu	Val	Gly	Trp	Arg	Trp	Ile

**RAW SEQUENCE LISTING**PATENT APPLICATION: US/09/788,188

DATE: 07/19/2001

TIME: 12:09:47

Input Set : A:\41673204.app

Output Set: N:\CRF3\07192001\1788188.raw

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294 225
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296 Arg Ile Asp Thr Ser Cys Val Cys Ala Leu Ser Arg Lys Ile Gly Arg
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299 Thr
303 <210> SEQ ID NO: 6
304 <211> LENGTH: 257
305 <212> TYPE: PRT
306 <213> ORGANISM: Artificial Sequence
308 <220> FEATURE:
309 <223> OTHER INFORMATION: Description of Artificial Sequence: Mutant NGF
          pro-neurotrophin
312 <400> SEQUENCE: 6
313 Met Ser Ile Leu Phe Tyr Val Ile Phe Leu Ala Tyr Leu Arg Gly Ile
316 Gln Gly Asn Asn Met Asp Gln Arg Ser Leu Pro Glu Asp Ser Leu Asn
317
                 20
                                      25
319 Ser Leu Ile Ile Lys Leu Ile Gln Ala Asp Ile Leu Lys Asn Lys Leu
322 Ser Lys Gln Met Val Asp Val Lys Glu Asn Tyr Gln Ser Thr Leu Pro
         50
325 Lys Ala Glu Ala Pro Arg Glu Pro Glu Arg Gly Gly Pro Ala Lys Ser
326
328 Ala Phe Gln Pro Val Ile Ala Met Asp Thr Glu Leu Leu Arg Gln Gln
329
                                          90
331 Arg Arg Tyr Asn Ser Pro Arg Val Leu Leu Ser Asp Ser Thr Pro Leu
                100
                                     105
334 Glu Pro Pro Pro Leu Tyr Leu Met Glu Asp Tyr Val Gly Ser Pro Val
335
            115
                                 120
337 Val Ala Ser Arg Thr Ser Arg Arg Lys Arg Tyr Ala Glu His Lys Ser
        130
                            135
340 His Arg Gly Glu Tyr Ser Val Cys Asp Ser Glu Ser Leu Trp Val Thr
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343 Asp Lys Ser Ser Ala Ile Asp Ile Arg Gly His Gln Val Thr Val Leu
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                                         170
346 Gly Glu Ile Lys Thr Gly Asn Ser Pro Val Lys Gln Tyr Phe Tyr Glu
347
                                     185
349 Thr Arg Cys Lys Glu Ala Arg Pro Val Lys Asn Gly Cys Arg Gly Ile
350
            195
                                 200
352 Asp Asp Lys His Trp Asn Ser Gln Cys Lys Thr Ser Gln Thr Tyr Val
                            215
                                                 220
355 Arg Ala Leu Thr Ser Glu Asn Asn Lys Leu Val Gly Trp Arg Trp Ile
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                                             235
358 Arg Ile Asp Thr Ser Cys Val Cys Ala Leu Ser Arg Lys Ile Gly Arg
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365 <210> SEQ ID NO: 7
366 <211> LENGTH: 210
367 <212> TYPE: PRT
368 <213> ORGANISM: Homo sapiens
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Use of n and/or Xaa has been detected in the Sequence Listing. Review the Sequence Listing to insure a corresponding explanation is presented in the <220> to <223> fields of each sequence using n or Xaa.

VERIFICATION SUMMARY

PATENT APPLICATION: US/09/788,188

DATE: 07/19/2001

TIME: 12:09:48

Input Set : A:\41673204.app

Output Set: N:\CRF3\07192001\1788188.raw

L:535 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:10 L:537 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:10